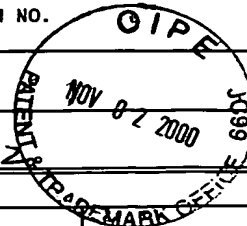


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FORM PTO-1449 INFORMATION DISCLOSURE STATEMENT	ATTY. DOCKET NO. 0609.4830001	APPLICATION NO. 09/617,116
	APPLICANT Aghi <i>et al.</i>	
	FILING DATE July 14, 2000	GROUP 1634 1632



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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB-CLASS	FILING DATE
QN	AA1	5,529,774	06/25/1996	Barba <i>et al.</i>	424	93.21	08/13/1991
	AB1	5,591,624	01/07/1997	Barber <i>et al.</i>	435	240.2	11/23/1993
	AC1	5,601,818	02/11/1997	Freeman <i>et al.</i>	424	93.21	12/08/1994
	AD1	5,631,236	05/20/1997	Woo <i>et al.</i>	514	44	08/26/1993
	AE1	5,670,488	09/23/1997	Gregory <i>et al.</i>	514	44	10/13/1993
	AF1	5,688,773	11/18/1997	Chiocca <i>et al.</i>	514	44	10/28/1994
	AG1	5,691,177	11/25/1997	Guber <i>et al.</i>	435	172.3	06/05/1995
	AH1	5,741,486	04/21/1998	Pathak <i>et al.</i>	424	93.21	06/06/1995
	AI1	5,763,217	06/09/1998	Cynader <i>et al.</i>	435	69.1	10/11/1995
QN	AJ1	5,763,242	06/09/1998	Zhang <i>et al.</i>	435	172.3	02/08/1995
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
QN	AL1	WO 95/06486	03/09/1995	WIPO			Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
QN	AR	1	Abdallah, B. <i>et al.</i> , "Non-viral gene transfer: Applications in developmental biology and gene therapy," <i>Biol. Cell</i> 85:1-7 (1995).
QN	AS	1	Aghi, M. <i>et al.</i> , "Synergistic Anticancer Effects of Ganciclovir/Thymidine Kinase and 5-Fluorocytosine/Cytosine Deaminase Gene Therapies," <i>J. Natl. Cancer Inst.</i> 90:370-380 (March 1998).
QN	AT	1	Aghi, M. <i>et al.</i> , "Folypolyglutamyl Synthetase Gene Transfer and Glioma Antifolate Sensitivity in Culture and <i>In Vivo</i> ," <i>J. Natl. Cancer Inst.</i> 91:1233-1241 (July 1999).

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				APPLICANT Aghi et al.			
				FILING DATE July 14, 2000		GROUP 1614	

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	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
QN	AR	2	Aghi, M. et al., "Prodrug activation enzymes in cancer gene therapy," <i>J. Gene Med.</i> 2:148-164 (May-June 2000).
QN	AS	2	Barredo, J. and Moran, R.G., "Determinants of Antifolate Cytotoxicity: Folylpolyglutamate Synthetase Activity during Cellular Proliferation and Development," <i>Molec. Pharmacol.</i> 42:687-694 (1992).
QN	AT	2	Calos, M.P. et al., "The potential of extrachromosomal replicating vectors for gene therapy," <i>Trends in Genetics</i> 12:463-467 (1996).

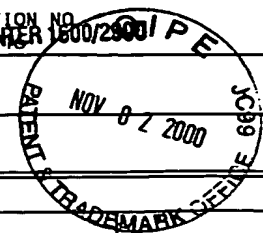
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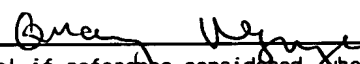
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FORM PTO-1449 <u>INFORMATION DISCLOSURE STATEMENT</u>				ATTY. DOCKET NO. 0609.4830001		APPLICATION NO. 0609.4830001	
				APPLICANT Aghi et al.			
				FILING DATE July 14, 2000			

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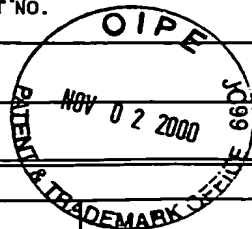
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	AR	3	Chabner, B.A. et al., "Polyglutamation of Methotrexate," <i>J. Clin. Invest.</i> 76:907-912 (1985)
	AS	3	Chase, M. et al., "An oncolytic viral mutant that delivers the CYP2B1 transgene and augments cyclophosphamide chemotherapy," <i>Nature Biotechnol.</i> 16:444-448 (1998).
 	AT	3	Chen, L. and Waxman, D.J., "Intratumoral Activation and Enhanced Chemotherapeutic Effect of Oxazaphosphorines following Cytochrome P-450 Gene Transfer: Development of a Combined Chemotherapy/Cancer Gene Therapy Strategy," <i>Cancer Res.</i> 55:581-589 (1995).

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0609.4830001APPLICANT
Aghi et al.FILING DATE
July 14, 2000GROUP
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	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

QW	AR	4	Chu, E. and Allegra, C.J., "Antifolates," in <i>Cancer Chemotherapy and Biotherapy: Principles and Practice</i> , 2 nd Edition, Chabner, B.A. and Longo, D.L., eds., Lippincott-Raven Publishers, New York, NY, pp. 109-148 (1996).
↓	AS	4	Dachs, G.U. et al., "Targeting Gene Therapy to Cancer: A Review," <i>Oncol. Res.</i> 9:313-325 (1997).
↓	AT	4	Danks, M.K. et al., "Overexpression of a Rabbit Liver Carboxylesterase Sensitizes Human Tumor Cells to CPT-11," <i>Cancer Res.</i> 58:20-22 (January 1998).

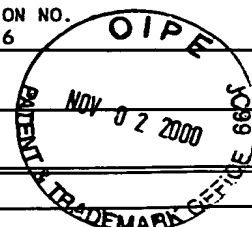
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AN	AR	5	Deonarain, M.P. et al., "Genetic Delivery of Enzymes for Cancer Therapy," <i>Gene Therapy</i> 2: 235-244 (1995).
I	AS	5	Drinkard, L. et al., "A Phase II Trial of Edatrexate (ETX) in Progressive Malignant Glioma (PMG)," <i>Program/Proceedings of the American Soc. of Clinical Oncology</i> 13:182, Abstract No. 517 (1994).
✓	AT	5	Freeman, S.M. et al., "The 'Bystander Effect': Tumor Regression When a Fraction of the Tumor Mass Is Genetically Modified," <i>Cancer Res.</i> 53:5274-5283 (1993).

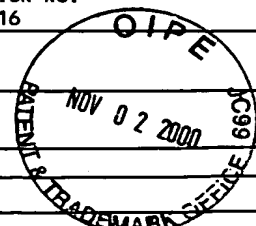
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AW	AR	6	Freeman, S.M. <i>et al.</i> , "In Situ Use of Suicide Genes for Cancer Therapy," <i>Semin. Oncology</i> 23:31-45 (1996).
J	AS	6	Freemantle, S.J. and Moran, R.G., "Transcription of the Human Polyglutamate Synthetase Gene," <i>J. Biol. Chem.</i> 272:25373-25379 (1997).
↓	AT	6	Fry, D.W. <i>et al.</i> , "Analysis of the Role of Membrane Transport and Polyglutamation of Methotrexate in Gut and the Ehrlich Tumor <i>in Vivo</i> as Factors in Drug Sensitivity and Selectivity," <i>Cancer Res.</i> 43:1087-1092 (1983).

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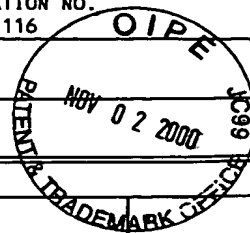
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

QW	AR	Z	Garrow, T.A. <i>et al.</i> , "Expression cloning of a human cDNA encoding folylpoly(γ -glutamate) synthetase and determination of its primary structure," <i>Proc. Natl. Acad. Sci. USA</i> 89:9151-9155 (1992).
I	AS	Z	Kim, J.-S. <i>et al.</i> , "Regulation of Folate and One-carbon Metabolism in Mammalian Cells," <i>J. Biol. Chem.</i> 268:21680-21685 (1993).
V	AT	Z	Lan, K-H. <i>et al.</i> , "In Vivo Selective Gene Expression and Therapy Mediated by Adenoviral Vectors for Human Carcinoembryonic Antigen-producing Gastric Carcinoma," <i>Cancer Res.</i> 57:4279-4284 (1997).

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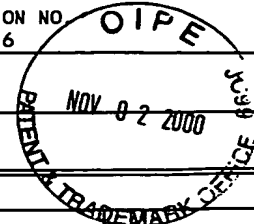
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

QW	AR	8	Li, W.-W. <i>et al.</i> , "Mechanisms of Natural Resistance to Antifolates in Human Soft Tissue Sarcomas," <i>Cancer Res.</i> 52:1434-1438 (1992).
↓	AS	8	Matherly, L.H. <i>et al.</i> , "Antifolate Polyglutamylolation and Competitive Drug Displacement at Dihydrofolate Reductase as Important Elements in Leucovorin Rescue in L1210 Cells," <i>Cancer Res.</i> 46:588-593 (1986).
↓	AT	8	McGuire, J.J. and Russell, C.A., "Folypolyglutamate Synthetase Expression in Antifolate-Sensitive and -Resistant Human Cell Lines," <i>Oncology Res.</i> 10:193-200 (1998).

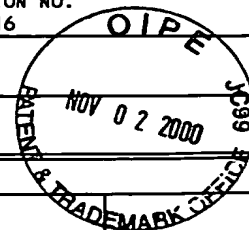
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	AL						Yes No
	AM						Yes No
	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

QW	AR	2	Miller, N. and Whelan, J., "Progress in Transcriptionally Targeted and Regulatable Vectors for Genetic Therapy," <i>Hum. Gene Ther.</i> 8:803-815 (1997).
↓	AS	2	Moolten, F.L., "Drug sensitivity ("suicide") genes for selective cancer chemotherapy," <i>Cancer Gene Ther.</i> 1:279-287 (1994).
↓	AT	2	Murdoch, B. <i>et al.</i> , "A rapid screening procedure for the identification of high-titer retrovirus packaging clones," <i>Gene Ther.</i> 4:744-749 (1997).

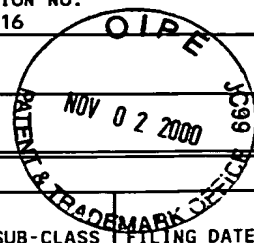
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

QW	AR	10	Nakanishi, M. <i>et al.</i> , "Gene Introduction Into Animal Tissues," <i>Crit. Rev. Therapeut. Drug Carrier Systems</i> 12:263-310 (1995).
J	AS	10	O'Malley, Jr., B.W. <i>et al.</i> , "Combination Gene Therapy for Oral Cancer in a Murine Model," <i>Cancer Res.</i> 56:1737-1741 (1996).
✓	AT	10	O'Malley, Jr., B.W. and Li, D., "Combination Gene Therapy for Salivary Gland Cancer," <i>Ann. N.Y. Acad. Sci.</i> 842:163-170 (April 1998).

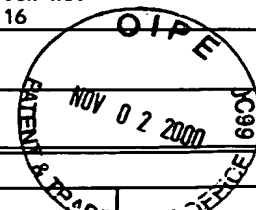
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

QW	AR	11	Peters, G.J. and van der Wilt, C.L., "Development of a Simple Folylpolyglutamate Synthetase Assay in Tissues and Cell Lines," in <i>Chemistry and biology of Pteridines and Folates</i> , Ayling, J.E. <i>et al.</i> , eds., Plenum Press, NY, pp. 651-654 (1993).
↓	AS	11	Pizzorno, G. <i>et al.</i> , "Impaired Polyglutamylation of Methotrexate as a Cause of Resistance in CCRF-CEM Cells after Short-Term, High-Dose Treatment with This Drug," <i>Cancer Res.</i> 48:2149-2155 (1988).
↓	AT	11	Rainov, N.G. <i>et al.</i> , "Retrovirus-Mediated Gene Therapy of Experimental Brain Neoplasms Using the Herpes Simplex Virus-Thymidine Kinase/Ganciclovir Paradigm," <i>Cancer Gene Therapy</i> 3:99-106 (1996).

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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AW	AR	12	Ram, Z. <i>et al.</i> , "In Situ Retroviral-mediated Gene Transfer for the Treatment of Brain Tumors in Rats," <i>Cancer Res.</i> 53:83-88 (1993).
V	AS	12	Rancourt, C. <i>et al.</i> , "Endothelial Cell Vehicles for Delivery of Cytotoxic Genes as a Gene Therapy Approach for Carcinoma of the Ovary," <i>Clin. Cancer Res.</i> 4:265-270 (February 1998).
V	AT	12	Robbins, P.D. <i>et al.</i> , "Viral vectors for gene therapy," <i>TIBTECH</i> 16:35-40 (January 1998).

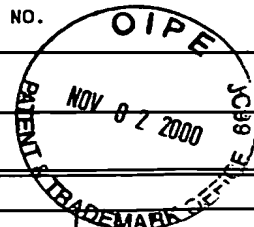
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	AP						Yes No

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aw	AR	13	Rogulski, K.R. <i>et al.</i> , "Glioma Cells Transduced with an <i>Escherichia coli</i> CD/HSV-1 TK Fusion Gene Exhibit Enhanced Metabolic Suicide and Radiosensitivity," <i>Human Gene Ther.</i> 8:73-85 (1997).
J	AS	13	Roth, J.A. <i>et al.</i> , "Retrovirus-Mediated Wild-Type p53 Gene Transfer to Tumors of Patients with Lung Cancer," <i>Nature Med.</i> 2:985-991 (1996).
J	AT	13	Roth, J.A. and Cristiano, R.J., "Gene Therapy for Cancer: What Have We Done and Where are we Going?" <i>J. Natl. Cancer Inst.</i> 89:21-39 (1997).

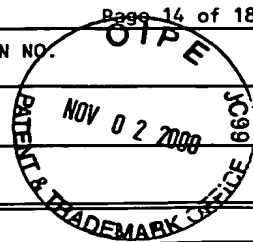
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FORM PTO-1449

INFORMATION DISCLOSURE STATEMENT

ATTY. DOCKET NO.
0609.4830001APPLICATION NO.
09/617,116APPLICANT
Aghi *et al.*FILING DATE
July 14, 2000GROUP
1614

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QW	AR	14	Roy, K. <i>et al.</i> , "Different Antifolate-resistant L1210 Cell Variants with either Increased or Decreased Folylpolyglutamate Synthetase Gene Expression at the Level of mRNA Transcription," <i>J. Biol. Chem.</i> 270:26918-26922 (1995).
J	AS	14	Roy, K. <i>et al.</i> , "Posttranscriptionally Mediated Decreases in Folylpolyglutamate Synthetase Gene Expression in Some Folate Analogue-resistant Variants of the L1210 Cell," <i>J. Biol. Chem.</i> 272:6903-6908 (1997).
✓	AT	14	Rumberger, B.G. <i>et al.</i> , "Differing Specificities for 4-Aminofolate Analogues of Folylpolyglutamyl Synthetase from Tumors and Proliferative Intestinal Epithelium of the Mouse with Significance for Selective Antitumor Action," <i>Cancer Res.</i> 50:4639-4643 (1990).

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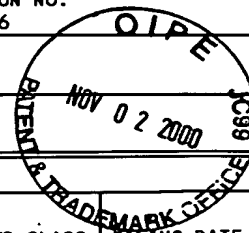
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OW	AR	15	Samuels, L.L. <i>et al.</i> , "Similar Differential for Total Polyglutamylation and Cytotoxicity among Various Folate Analogues in Human and Murine Tumor Cells <i>in Vitro</i> ," <i>Cancer Res.</i> 45:1488-1495 (1985).
	AS	15	Takemura, Y. and Jackman, A.L., "Folate-based thymidylate synthase inhibitors in cancer chemotherapy," <i>Anti-Cancer drugs</i> 8:3-16 (1997).
	AT	15	Takemura, Y. <i>et al.</i> , "Altered Expression of Folylpolyglutamate Synthetase (FPGS) Gene in Human Leukaemia Cells with Defective Polyglutamation of Raltitrexed (ZD1694)," <i>Brit. J. Cancer</i> 75(suppl. 1):31, Abstract No. P52 (1997).

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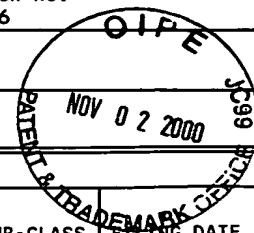
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	AR	16	Taylor, S.M. <i>et al.</i> , "Structural Organization of the Human Folylpoly-γ-glutamate Synthetase Gene: Evidence for a Single Genomic Locus," <i>Cancer Res.</i> 55: 6030-6034 (1995).
	AS	16	Terzis, A.J.A. <i>et al.</i> , "Proliferation, Migration and Invasion of Human Glioma Cells Exposed to Antifolate Drugs," <i>Int. J. Cancer</i> 54:112-118 (1993).
	AT	16	Uckert, W. <i>et al.</i> , "Double Suicide Gene (Cytosine Deaminase and Herpes Simplex Virus Thymidine Kinase) but Not Single Gene Transfer Allows Reliable Elimination of Tumor Cells <i>In Vivo</i> ," <i>Human Gene Ther.</i> 9:855-865 (April 1998).

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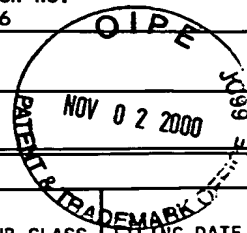
	AR	17	Wei, M.X. <i>et al.</i> , "Experimental Tumor Therapy in mice Using the Cyclophosphamide-Activating Cytochrome P450 2B1 Gene," <i>Hum. Gene Ther.</i> 5:969-978 (1994).
	AS	17	Yu, J.S. <i>et al.</i> , "Retroviral Delivery and Tetracycline-dependent Expression of IL-1 β -converting Enzyme (ICE) in a Rat Glioma Model Provides Controlled Induction of Apoptotic Death in Tumor Cells," <i>Cancer Res.</i> 56:5423-5427 (1996).
	AT	17	Zhang, J. and Russell, S.J., "Vectors for cancer gene therapy," <i>Cancer & Metastasis Rev.</i> 15:385-401 (1996).

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QW	AR	18	Zufferey, R. <i>et al.</i> , "Multiply attenuated lentiviral vector achieves efficient gene delivery in vivo," <i>Nature Biotechnol.</i> 15:871-875 (1997).
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